

CLAIMS

1. HIV-2 retrovirus or variance of this virus, which retrovirus has infectious properties with respect to human T4 lymphocytes and the essential morphological and immunological properties of any of the retroviruses deposited at the CNCM under n° I-502, I-532, I-642 and I-643.

2. The purified retrovirus of claim 1 which possesses the following properties :

- 10 - the preferred target for the HIV-2 retrovirus consists of human Leu 3 cells (or T4 lymphocytes) and for permanent cell lines derived of said T4 lymphocytes ;
- it is cytotoxic for the human T4 lymphocytes which it infects ;
- 15 - it has a reverse transcriptase activity which requires the presence of Mg^{2+} ions and has a strong affinity for poly adenylate oligodeoxythymidylate (poly(A)-oligo(dT) 12-18) ;
- it has a density of approximately 1.16 in a sucrose gradient ;
- 20 - it has a mean diameter of 140 nanometres and a core having mean diameter of 41 nanometres ;
- it can be cultivated in permanent cell lines expressing the T4 protein ;
- 25 - it is not infectious in T8 lymphocytes ;
- the lysates of this virus contain p26 protein which does not crossreact immunologically with p24 protein of the HTLV-1 virus or of the HTLV-2 ;
- said lysates further contain p-16 protein which is not recognized immunologically by p19 protein of HTLV-1 or
- 30 of HTLV-2 in radioimmunoprecipitation assays ;
- said lysates further contain an envelope glycoprotein having a molecular weight of the order of 130,000-140,000 which does not crossreact immunologically with

gp110 of HTLV-1 retrovirus ;

- said lysates further contain a molecule which can be labelled by ³⁵S-cystein, having an apparent molecular weight of about 36,000 ;

5 - the genomic RNA of HIV-2 hybridizes neither with the genomic RNA, nor with the ENV gene, nor with the LTRs of HIV-1 under stringent conditions ;

10 - the genomic RNA of HIV-2 hybridizes weakly under non-stringent conditions with nucleotide sequences of the CAG region of the HIV-1 genome.

3. The retrovirus of claim 2 whose lysates also contain a molecule having an apparent molecular weight of 42,000-45,000

15 4. The retrovirus of any of claims 1 to 3, wherein the nucleotidic sequence of its genomic RNA which comprises the R region and the U3 region also comprises a nucleotidic sequence which corresponds with the following nucleotide sequence :

20 GTGGAAGGCGAGACTGAAAGCAAGAGGAATACCATTTAGTTAAAGGACAG
GAACAGCTATACTTGCTCAGGGCAGGAAGTAACTAACAGAAACAGCTGAG
ACTGCAGGGACTTTCCAGAAGGGGCTGTAACCAAGGGAGGGACATGGGAG
GAGCTGGTGGGGAAACGCTCATATTCTCTGTATAATATACCCGCTGCTTG
25 CATTGTACTTCAGTCGCTCTGCGGAGAGGCTGGCAGATTGAGCCCTGGAG
GATCTCTCCAGCACTAGACGGATGAGCCTGGGTGCCCTGCTAGACTCTCA
CCAGCACTTGGCCGGTGCTGGCAGACGGCCCCACGCTTGCTGCTTAAAA
ACCTTCCTTAATAAAGCTGCAGTAGAAGCA

5. The retrovirus of anyone of claims 1 to 4 whose genomic RNA also contains a GAG sequence which corresponds with the following nucleotide sequence :

GAGROD

5

ATGGGCGCGAGAACTCCGTCTTGAGAGGGGAAAAAAGCAGATGAA

TTAGAAAGAATCAGGTTACGGCCCGGCGAAAGAAAAAGTACAGG

10

CTAAACATATTGTGTGGGCAGCGAATAAATTGGACAGATTCCGA
100

TTAGCAGAGAGCCTGTTGGAGTCAAAGAGGGTTGTCAAAAAATT

15

CTTACAGTTTTAGATCCAATGGTACCGACAGGTTCAGAAAAATT
200

AAAAGTCTTTTTAATACTGTCTGCGTCATTTGGTGCATACAGCA

GAAGAGAAAGTGAAAGATACTGAAGGAGCAAAACAAATAGTGCGG
300

20

AGACATCTAGTGGCAGAAACAGGAACTGCAGAGAAAATGCCAAGC

ACAAGTAGACCAACAGCACCATCTAGCGAGAAGGGAGGAAATTAC
400

25

CCAGTGCAACATGTAGGCGGCAACTACACCCATATACCGCTGAGT

CCCCGAACCTAAATGCCCTGGGTAAAATTAGTAGAGGAAAAAAG

TTGGGGCAGAAGTAGTGCCAGGATTTAGGCACTCTCAGAAGGC
500

30

TGCACGCCCTATGATATCAACCAAATGCTTAATTGTGTGGGCGAC

CATCAAGCAGCCATGCAGATAATCAGGGAGATTATCAATGAGGAA
600

35

GCAGCAGAATGGGATGTGCAACATCCAATACCAGGCCCTTACCA

GCGGGGCAGCTTAGAGAGCCAAGGGGATCTGACATAGCAGGGACA
700

ACAAGCACAGTGAAGAACAGATCCAGTGGATGTTTAGGCCACAA

AATCCTGTACCAGTAGGAAACATCTATAGAAGATGGATCCAGATA
800

GGATTGCAGAAGTGTGTCAGGATGTACAACCCGACCAACATCCTA

GACATAAAACAGGGACCAAAGGAGCCGTTCCAAAGCTATGTAGAT
900

AGATTCTACAAAAGCTTGAGGGCAGAACAAACAGATCCAGCAGTG

AAGAATTGGATGACCCAAACACTGCTAGTACAAAATGCCAACCCA

GACTGTAAATTAGTGCTAAAAGGACTAGGGATGAACCCTACCTTA
1000

GAAGAGATGCTGACCGCCTGTGAGGGGTAGGTGGGCCAGGCCAG

AAAGCTAGATTAATGGCAGAGGCCCTGAAAGAGGTCATAGGACCT
1100

CCCCCTATCCCATTCGCAGCAGCCAGCAGAGAAAGGCATTTAAA

TGCTGGAAGTGTGAAAGGAAGGGCACTGGCAAGACAATGCCGA
1200

GCACCTAGAAGGCAGGGCTGCTGGAAGTGTGGTAAGCCAGGACAC

ATCATGACAAACTGCCCAGATAGACAGGCAGGTTTTTTAGGACTG
1300

GGCCCTTGGGGAAAGAAGCCCCGCAACTTCCCCGTGGCCCAAGTT

CCGCAGGGGCTGACACCAACAGCACCCCCAGTGGATCCAGCAGTG

GATCTACTGGAGAAATATATGCAGCAAGGGAAGACAGAGAGAG
1400

CAGAGAGAGAGACCATAAAGGAAGTGACAGAGGACTTACTGCAC

CTCGAGCAGGGGGAGACACCATAACAGGGAGCCACCAACAGAGGAC
1500

TTGCTGCACCTCAATTCTCTCTTTGGAAAAGACCAG

6. The retrovirus of anyone of claims 1 to 5 whose genomic RNA contains an ENV sequence which corresponds with the following nucleotide sequence :

ENVRN

5
ATGATGAATCAGCTGCTTATTGCCATTTTATTAGCTAGTGCTTGC
TTAGTATATTGCACCCAATATGTAAC TGTTTCTATGGCGTACCC
10 ACGTGGA AAAAATGCAACCATTCCCTCTTTTG TGCAACCAGAAAT
100
AGGGATACTTGGGGAACCATACAGTGCTTGCGCTGACAATGATGAT
TATCAGGAAATAACTTTGAATGTAACAGAGGCTTTTGATGCATGG
15 200
AATAATACAGTAACAGAACAAAGCAATAGAAGATGTCTGGCATCTA
TTCGAGACATCAATAAAACCATGTGTCAAAC TAACACCTTTATGT
300
20 GTAGCAATGAAATGCAGCAGCACAGAGAGCAGCACAGGGAACAAC
ACAACCTCAAAGAGCACAAGCACAACCACAACCACACCCACAGAC
400
25 CAGGAGCAAGAGATAAGTGAGGATACTCCATGCGCAGCGCAGAC
AACTGCTCAGGATTGGGAGAGGAAGAAACGATCAATTGCCAGTTC
AATATGACAGGATTAGAAAGAGATAAGAAAAACAGTATAATGAA
300
30 ACATGGTACTCAAAA GATGTGGTTTGTGAGACAAATAATAGCACA
AATCAGACCCAGTGTTACATGAACCATTGCAACACATCAGTCATC
600
35 ACAGAATCATGTGACAAGCACTATTGGGATGCTATAAGGTTTAGA
TACTGTGCACCACCGGGTTATGCCCTATTAAGATGTAATGATACC
700
AATTATT CAGGCTTTGCACCCA ACTGTTCTAAAGTAGTAGCTTCT

ACATGCACCAGGATGATGGAACGCAAACCTCCACATGGTTTGGC
800

TTAATGGCACTAGAGCAGAGAATAGAACATATATCTATTGGCAT

GGCAGAGATAATAGAACTATCATCAGCTTAAACAAATATTATAAT
900

CTCAGTTTGCATTGTAAGAGGCCAGGGAATAAGACAGTGAAACAA

ATAATGCTTATGTCAGGACATGTGTTTCACTCCCACTACCAGCCG

ATCAATAAAAGACCCAGACAAGCATGGTGCTGGTTCAAAGGCAAA
1000

TGGAAGACGCCATGCAGGAGGTGAAGACCCTTGCAAAACATCCC

AGGTATAGAGGAACCAATGACACAAGGAATATTAGCTTTGCAGCG
1100

CCAGGAAAAGGCTCAGACCCAGAAGTAGCATACATGTGGACTAAC

TGCAGAGGAGAGTTTCTCTACTGCAACATGACTTGGTTCTCAAT
1200

TGGATAGAGAATAAGACACACCGCAATTATGCACCGTGCCATATA

AAGCAAATAATTAACACATGGCATAAGGTAGGGAGAAATGTATAT
1300

TTGCCTCCCAGGGAAGGGGAGCTGTCTTGCAACTCAACAGTAACC

AGCATAATTGCTAACATTGACTGGCAAAACAATAATCAGACAAAC

ATTACCTTTAGTGGAGAGGTGGCAGAACTATACAGATTGGAGTTG
1400

GGAGATTATAAATTGGTAGAAATAACACCAATTGGCTTCGCACCT

ACAAAAGAAAAAAGATACTCCTCTGCTCACGGGAGACATACAAGA
1500
GGTGTGTTTCGTGCTAGGGTTCTTGGGTTTTCTCGCAACAGCAGGT
TCTGCAATGGGCGCTCGAGCGTCCCTGACCGTGTGGGCTCAGTCC
1600
CGGACTTTACTGGCCGGGATAGTGCAGCAACAGCAACAGCTGTTG
GACGTGGTCAAGAGACAACAAGAACTGTTGCGAGTGACCGTCTGG
1700
GGAACGAAAAACCTCCAGGCAAGAGTCACTGCTATAGAGAAGTAC
CTACAGGACCAGGCGCGCTAAATTCATGGGGATGTGCGTTTAGA
1800
CAAGTCTGCCACACTACTGTACCATGGGTTAATGATTCTTAGCA
CCTGACTGGGACAATATGACGTGGCAGGAATGGGAAAAACAAGTC
CGCTACCTGGAGGCAAATATCAGTAAAAGTTAGAACAGGCACAA
1900
ATTCAGCAAGAGAAAAATATGTATGAACTACAAAAATTAAATAGC
TGGGATATTTTTGGCAATTGGTTTGACTTAACCTCCTGGGTCAAG
2000
TATATTCAATATGGAGTGCTTATAATAGTAGCAGTAATAGCTTTA
AGAATAGTGATATATGTAGTACAAATGTTAAGTAGGCTTAGAAAG
2100
GGCTATAGGCCCTGTTTTCTCTTCCCCCCCCGGTTATATCCAACAG

ATCCATATCCACAAGGACCGGGAGAGCCAGCCAACGAAGAAACA
2200

GAAGAAGACGGTGGAGCAACGGTGGAGACAGATACTGGCCCTGG

CCGATAGCATATATACATTTCTGATCCGCCAGCTGATTGCGCTC

TTGACCAGACTATACAGCATCTGCAGGACTTACTATCCAGGAGC
2300

TTCTGACCTTCCAACATCTACCAGAATCTCAGAGACTGGCTG

AGACTTAGAACAGCCTTCTTGCAATATGGGTGCCAGTGGATCAA
2400

GAAGCATTCCAGGCCGCCGCGAGGGCTACAAGAGAGACTCTTGCG

GGCGCGTGCAGGGGCTTGTGGAGGGTATTGGAACGAATCGGGAGG
2500

GGAATACTCGCGGTTCCAAGAAGGATCAGACAGGGAGCAGAAATC

GCCCTCCTGTGAGGGACGGCAGTATCAGCAGGGAGACTTTATGAA
2600

TACTCCATGGAAGGACCCAGCAGCAGAAAGGGAGAAAAATTTGTA

CAGGCAACAAATATGGA

7. The retrovirus of anyone of claims 1 to 6 whose RNA virtually hybridizes neither with the ENV gene and the LTR close to it, particularly with the nucleotide sequence 5290-9130 of HIV-1, nor with the sequences of the POL region of the HIV-1 genome, particularly with the nucleotide sequence 2170-2240 of HIV-1.

8. A composition comprising at least one antigen, particularly a protein or glycoprotein of HIV-2 virus according to anyone of claims 1 to 7.

9. The composition of claim 8 which consists of total extract or lysate of said retrovirus.

10. The composition of claim 8 wherein said antigen consists of at least one of the internal core proteins of said virus, particularly p12, p16 and p26, which have apparent molecular weight of the order of 12,000, 16,000 and 26,000.

11. The composition of claim 8, characterized in that it contains a gp140 glycoprotein having an apparent molecular weight of about 130,000-140,000.

12. An antigen which provides a single band in electrophoresis on a polyacrylamid gel which comprises, in common with one of the purified antigens of HIV-2 retrovirus, an epitope that is recognized by the serum of a carrier of antibody against HIV-2.

13. A purified antigen having the immunological characteristics of one of the following proteins or glycoproteins of HIV-2: p12, p16, p26, p36, p42 and gp140.

14. An antigen of claim 13 which has the following aminoacid sequence or a part of said sequence recognized by anti-p12 antibodies :

Arg Lys Ala Phe Lys
 Cys Trp Asn Cys Gly Lys Glu Gly His Ser Ala Arg Gln Cys Arg
 1200
 Ala Pro Arg Arg Gln Gly Cys Trp Lys Cys Gly Lys Pro Gly His
 Ile Met Thr Asn Cys Pro Asp Arg Gln Ala Gly Phe Leu Gly Leu
 1300
 Gly Pro Trp Gly Lys Lys Pro Arg Asn Phe Pro Val Ala Gln Val
 Pro Gln Gly Leu Thr Pro Thr Ala Pro Pro Val Asp Pro Ala Val
 Asp Leu Leu Glu Lys Tyr Met Gln Gln Gly Lys Arg Gln Arg Glu
 1400
 Gln Arg Glu Arg Pro Tyr Lys Glu Val Thr Glu Asp Leu Leu His
 Leu Glu Gln Gly Glu Thr Pro Tyr Arg Glu Pro Pro Thr Glu Asp
 1500
 Leu Leu His Leu Asn Ser Leu Phe Gly Lys Asp Gln

15. An antigen of claim 13 which has the following amino acid sequence or a part of said sequence recognized by anti-p16 antibodies :

5 MetGlyAlaArgAsnSerValLeuArgGlyLysLysAlaAspGlu
 LeuGluArgIleArgLeuArgProGlyGlyLysLysLysTyrArg
 LeuLysHisIleValTrpAlaAlaAsnLysLeuAspArgPheGly
 10 100
 LeuAlaGluSerLeuLeuGluSerLysGluGlyCysGluLysIle
 LeuThrValLeuAspProMetValProThrGlySerGluAsnLeu
 15 200
 LysSerLeuPheAsnThrValCysValIleTrpCysIleHisAla
 GluGluLysValLysAspThrGluGlyAlaLysGlnIleValArg
 300
 ArgHisLeuValAlaGluThrGlyThrAlaGluLysMetProSer
 20 ThrSerArgProThrAlaProSerSerGluLysGlyGlyAsnTyr
 400

16. An antigen of claim 13 which has the following aminoacid sequence or a part of said sequence recognized by anti-p26 antibodies :

5 ProValGlnHisValGlyGlyAsnTyrThrHisIleProLeuSer
 ProArgThrLeuAsnAlaTrpValLysLeuValGluGluLysLys
 PheGlyAlaGluValValProGlyPheGlnAlaLeuSerGluGly
 10 500
 CysThrProTyrAspIleAsnGlnMetLeuAsnCysValGlyAsp
 HisGlnAlaAlaMetGlnIleIleArgGluIleIleAsnGluGlu
 600
 15 AlaAlaGluTrpAspValGlnHisProIleProGlyProLeuPro
 AlaGlyGlnLeuArgGluProArgGlySerAspIleAlaGlyThr
 700
 ThrSerThrValGluGluGlnIleGlnTrpMetPheArgProGln
 20 AsnProValProValGlyAsnIleTyrArgArgTrpIleGlnIle
 800
 GlyLeuGlnLysCysValArgMetTyrAsnProThrAsnIleLeu
 AspIleLysGlnGlyProLysGluProPheGlnSerTyrValAsp
 25 ArgPheTyrLysSerLeuArgAlaGluGlnThrAspProAlaVal
 900
 LysAsnTrpMetThrGlnThrLeuLeuValGlnAsnAlaAsnPro
 30 AspCysLysLeuValLeuLysGlyLeuGlyMetAsnProThrLeu
 1000
 GluGluMetLeuThrAlaCysGlnGlyValGlyGlyProGlyGln
 LysAlaArgLeuMetAlaGluAlaLeuLysGluValIleGlyPro
 35 1100
 AlaProIleProPheAlaAlaAlaGlnGln

17. An antigen of claim 13 which has the following aminoacid sequence or a part of said sequence recognized by anti-gp140 antibodies :

ENVRN

5 MetMetAsnGlnLeuLeuIleAlaIleLeuLeuAlaSerAlaCys
 LeuValTyrCysThrGlnTyrValThrValPheTyrGlyValPro
 10 ThrTrpLysAsnAlaThrIleProLeuPheCysAlaThrArgAsn
 100 ArgAspThrTrpGlyThrIleGlnCysLeuProAspAsnAspAsp
 TyrGlnGluIleThrLeuAsnValThrGluAlaPheAspAlaTrp
 15 AsnAsnThrValThrGluGlnAlaIleGluAspValTrpHisLeu
 PheGluThrSerIleLysProCysValLysLeuThrProLeuCys
 20 ValAlaMetLysCysSerSerThrGluSerSerThrGlyAsnAsn
 ThrThrSerLysSerThrSerThrThrThrThrThrProThrAsp
 400 GlnGluGlnGluIleSerGluAspThrProCysAlaArgAlaAsp
 25 AsnCysSerGlyLeuGlyGluGluGluThrIleAsnCysGlnPhe
 AsnMetThrGlyLeuGluArgAspLysLysLysGlnTyrAsnGlu
 500 ThrTrpTyrSerLysAspValValCysGluThrAsnAsnSerThr
 30 AsnGlnThrGlnCysTyrMetAsnHisCysAsnThrSerValIle
 600 ThrGluSerCysAspLysHisTyrTrpAspAlaIleArgPheArg
 35 TyrCysAlaProProGlyTyrAlaLeuLeuArgCysAsnAspThr
 700 AsnTyrSerGlyPheAlaProAsnCysSerLysValValAlaSer

ThrCysThrArgMetMetGluThrGlnThrSerThrTrpPheGly
 PheAsnGlyThrArgAlaGluAsnArgThrTyrIleTyrTrpHis 800
 GlyArgAspAsnArgThrIleIleSerLeuAsnLysTyrTyrAsn
 LeuSerLeuHisCysLysArgProGlyAsnLysThrValLysGln 900
 IleMetLeuMetSerGlyHisValPheHisSerHisTyrGlnPro
 IleAsnLysArgProArgGlnAlaTrpCysTrpPheLysGlyLys
 TrpLysAspAlaMetGlnGluValLysThrLeuAlaLysHisPro 1000
 ArgTyrArgGlyThrAsnAspThrArgAsnIleSerPheAlaAla
 ProGlyLysGlySerAspProGluValAlaTyrMetTrpThrAsn 1100
 CysArgGlyGluPheLeuTyrCysAsnMetThrTrpPheLeuAsn
 TrpIleGluAsnLysThrHisArgAsnTyrAlaProCysHisIle 1200
 LysGlnIleIleAsnThrTrpHisLysValGlyArgAsnValTyr
 LeuProProArgGluGlyGluLeuSerCysAsnSerThrValThr 1300
 SerIleIleAlaAsnIleAspTrpGlnAsnAsnAsnGlnThrAsn
 IleThrPheSerAlaGluValAlaGluLeuTyrArgLeuGluLeu
 GlyAspTyrLysLeuValGluIleThrProIleGlyPheAlaPro 1400

ThrLysGluLysArgTyrSerSerAlaHisGlyArgHisThrArg
 1500
 GlyValPheValLeuGlyPheLeuGlyPheLeuAlaThrAlaGly
 SerAlaMetGlyAlaArgAlaSerLeuThrValSerAlaGlnSer
 1600
 ArgThrLeuLeuAlaGlyIleValGlnGlnGlnGlnGlnLeuLeu
 AspValValLysArgGlnGlnGluLeuLeuArgLeuThrValTrp
 1700
 GlyThrLysAsnLeuGlnAlaArgValThrAlaIleGluLysTyr
 LeuGlnAspGlnAlaArgLeuAsnSerTrpGlyCysAlaPheArg
 1800
 GlnValCysHisThrThrValProTrpValAsnAspSerLeuAla
 ProAspTrpAspAsnMetThrTrpGlnGluTrpGluLysGlnVal
 ArgTyrLeuGluAlaAsnIleSerLysSerLeuGluGlnAlaGln
 1900
 IleGlnGlnGluLysAsnMetTyrGluLeuGlnLysLeuAsnSer
 TrpAspIlePheGlyAsnTrpPheAspLeuThrSerTrpValLys
 2000
 TyrIleGlnTyrGlyValLeuIleIleValAlaValIleAlaLeu
 ArgIleValIleTyrValValGlnMetLeuSerArgLeuArgLys
 2100
 GlyTyrArgProValPheSerSerProProGlyTyrIleGlnGln

IleEisIleEisLysAspArgGlyGlnProAlaAsnGluGluThr
 GluGluAspGlyGlySerAsnGlyGlyAspArgTyrTrpProTrp 2200
 ProIleAlaTyrIleHisPheLeuIleArgGlnLeuIleArgLeu
 LeuThrArgLeuTyrSerIleCysArgAspLeuLeuSerArgSer
 2300
 PheLeuThrLeuGlnLeuIleTyrGlnAsnLeuArgAspTrpLeu
 ArgLeuArgThrAlaPheLeuGlnTyrGlyCysGluTrpIleGln
 2400
 GluAlaPheGlnAlaAlaAlaArgAlaThrArgGluThrLeuAla
 GlyAlaCysArgGlyLeuTrpArgValLeuGlnArgIleGlyArg
 2500
 GlyIleLeuAlaValProArgArgIleArgGlnGlyAlaGluIle
 AlaLeuLeu***GlyThrAlaValSerAlaGlyArgLeuTyrGlu
 2600
 TyrSerMetGluGlyProSerSerArgLysGlyGluLysPheVal
 GlnAlaThrLysTyrGly

18. A method for the in vitro detection of the presence of antibodies against anti-HIV-2 in a biological liquid, such as a serum, more particularly for the in vitro diagnosis of a potential or existing LAS or AIDS caused by HIV-2 type retrovirus, which comprises contacting a serum or other biological medium from the person to be diagnosed with a composition according to anyone of claims 8 to 11 or with an antigen according to anyone of claims 12 to 17, detecting the immunological conjugate possibly formed between said anti-HIV-2-antibodies and the antigen or antigens used.

19. The method of claim 18 which comprises achieving the detection of said immunological conjugate by reacting said immunological conjugate possibly formed with a labelled reagent formed either by human anti-immunoglobulin-antibodies or of a bacterial A protein, and by detecting the complexe formed between the reagent and said immunological conjugate.

20. Kit for the detection of anti-HIV-2-antibodies in a biological fluid, particularly of a person possibly carrying such antibodies, which comprises :

- a composition such as defined in anyone of claims 8 to 11 or an antigen such as defined in any of claims 12 to 17 ; and
- means for detecting the immunological complexe resulting from the immunological reaction between the antigen and said biological fluid.

21. The kit of claim 21, whose means for detecting the immunological complexe formed comprises human anti-immunoglobulins or a protein A and a means for detecting the complexe formed between the anti-HIV-2 antibodies contained in the detected immunological conjugate.

22. Immunogenic compositions containing an envelope glycoprotein of HIV-2 retrovirus, such as gp140

of said retrovirus, or part of said glycoprotein, in association with a pharmaceutically acceptable vehicle appropriate for the constitution of vaccines effective against HIV-2.

- 5 23. The composition of claim 22 which contains at least part of an immunogenic glycoprotein comprising the proteic backbone having the following sequence :

ENVRN

MetMetAsnGlnLeuLeuIleAlaIleLeuLeuAlaSerAlaCys
 LeuValTyrCysThrGlnTyrValThrValPheTyrGlyValPro
 ThrTrpLysAsnAlaThrIleProLeuPheCysAlaThrArgAsn
 100
 ArgAspThrTrpGlyThrIleGlnCysLeuProAspAsnAspAsp
 TyrGlnGluIleThrLeuAsnValThrGluAlaPheAspAlaTrp
 200
 AsnAsnThrValThrGluGlnAlaIleGluAspValTrpHisLeu
 PheGluThrSerIleLysProCysValLysLeuThrProLeuCys
 300
 ValAlaMetLysCysSerSerThrGluSerSerThrGlyAsnAsn
 ThrThrSerLysSerThrSerThrThrThrThrProThrAsp
 400
 GlnGluGlnGluIleSerGluAspThrProCysAlaArgAlaAsp
 AsnCysSerGlyLeuGlyGluGluGluThrIleAsnCysGlnPhe
 AsnMetThrGlyLeuGluArgAspLysLysLysGlnTyrAsnGlu
 500
 ThrTrpTyrSerLysAspValValCysGluThrAsnAsnSerThr
 AsnGlnThrGlnCysTyrMetAsnHisCysAsnThrSerValIle
 600
 ThrGluSerCysAspLysHisTyrTrpAspAlaIleArgPheArg
 TyrCysAlaProProGlyTyrAlaLeuLeuArgCysAsnAspThr
 700
 AsnTyrSerGlyPheAlaProAsnCysSerLysValValAlaSer

ThrCysThrArgMetMetGluThrGlnThrSerThrTrpPheGly
 PheAsnGlyThrArgAlaGluAsnArgThrTyrIleTyrTrpHis 800
 GlyArgAspAsnArgThrIleIleSerLeuAsnLysTyrTyrAsn
 LeuSerLeuHisCysLysArgProGlyAsnLysThrValLysGln 900
 IleMetLeuMetSerGlyHisValPheHisSerHisTyrGlnPro
 IleAsnLysArgProArgGlnAlaTrpCysTrpPheLysGlyLys
 TrpLysAspAlaMetGlnGluValLysThrLeuAlaLysHisPro 1000
 ArgTyrArgGlyThrAsnAspThrArgAsnIleSerPheAlaAla
 ProGlyLysGlySerAspProGluValAlaTyrMetTrpThrAsn 1100
 CysArgGlyGluPheLeuTyrCysAsnMetThrTrpPheLeuAsn
 TrpIleGluAsnLysThrHisArgAsnTyrAlaProCysHisIle 1200
 LysGlnIleIleAsnThrTrpHisLysValGlyArgAsnValTyr
 LeuProProArgGluGlyGluLeuSerCysAsnSerThrValThr 1300
 SerIleIleAlaAsnIleAspTrpGlnAsnAsnAsnGlnThrAsn
 IleThrPheSerAlaGluValAlaGluLeuTyrArgLeuGluLeu
 GlyAspTyrLysLeuValGluIleThrProIleGlyPheAlaPro 1400

ThrLysGluLysArgTyrSerSerAlaHisGlyArgHisThrArg

1500

GlyValPheValLeuGlyPheLeuGlyPheLeuAlaThrAlaGly

SerAlaMetGlyAlaArgAlaSerLeuThrValSerAlaGlnSer

1600

ArgThrLeuLeuAlaGlyIleValGlnGlnGlnGlnGlnLeuLeu

AspValValLysArgGlnGlnGluLeuLeuArgLeuThrValTrp

1700

GlyThrLysAsnLeuGlnAlaArgValThrAlaIleGluLysTyr

LeuGlnAspGlnAlaArgLeuAsnSerTrpGlyCysAlaPheArg

1800

GlnValCysHisThrThrValProTrpValAsnAspSerLeuAla

ProAspTrpAspAsnMetThrTrpGlnGluTrpGluLysGlnVal

ArgTyrLeuGluAlaAsnIleSerLysSerLeuGluGlnAlaGln

1900

IleGlnGlnGluLysAsnMetTyrGluLeuGlnLysLeuAsnSer

TrpAspIlePheGlyAsnTrpPheAspLeuThrSerTrpValLys

2000

TyrIleGlnTyrGlyValLeuIleIleValAlaValIleAlaLeu

ArgIleValIleTyrValValGlnMetLeuSerArgLeuArgLys

2100

GlyTyrArgProValPheSerSerProProGlyTyrIleGlnGln

IleEisIleEisLysAspArgGlyGlnProAlaAsnGluGluThr
 GluGluAspGlyGlySerAsnGlyGlyAspArgTyrTrpProTrp 2200
 ProIleAlaTyrIleHisPheLeuIleArgGlnLeuIleArgLeu
 LeuThrArgLeuTyrSerIleCysArgAspLeuLeuSerArgSer
 2300
 PheLeuThrLeuGlnLeuIleTyrGlnAsnLeuArgAspTrpLeu
 ArgLeuArgThrAlaPheLeuGlnTyrGlyCysGluTrpIleGln
 2400
 GluAlaPheGlnAlaAlaAlaArgAlaThrArgGluThrLeuAla
 GlyAlaCysArgGlyLeuTrpArgValLeuGluArgIleGlyArg
 2500
 GlyIleLeuAlaValProArgArgIleArgGlnGlyAlaGluIle
 AlaLeuLeu***GlyThrAlaValSerAlaGlyArgLeuTyrGlu
 2600
 TyrSerMetGluGlyProSerSerArgLysGlyGluLysPheVal
 GlnAlaThrLysTyrGly

24. The immunogenic composition of claim 22 or of claim 23 which is dosed in antigen in order to enable the administration of a dosage-unit of 10 to 500, particularly from 50 to 100 µg/kg of bodyweight.

5 25. Monoclonal antibody characterized by its ability to specifically recognize one of the antigens according to anyone of claims 14 to 17.

26. The secreting hybridomas of the monoclonal antibody of claim 25.

10 27. Nucleic acids, optionally labelled, derived of part at least of RNA of HIV-2 virus or of one of its variance.

15 28. The nucleic acid of claim 27, which contains at least part of the cDNA which corresponds with the entire genomic RNA of HIV-2 retrovirus.

29. The nucleic acid of claim 27, which contains the nucleotide sequence :

20 GTGGAAGGCGAGACTGAAAGCAAGAGGAATACCATTTAGTTAAAGGACAG
GAACAGCTATACTTGGTCAGGGCAGGAAGTAACTAACAGAAACAGCTGAG
ACTGCAGGGACTTTCCAGAAGGGGCTGTAACCAAGGGAGGGACATGGGAG
GAGCTGGTGGGGAACGCCTCATATTCTCTGTATAATATACCCGCTGCTTG
CATTGTACTTCAGTCGCTCTGCCGAGAGGCTGGCAGATTGAGCCCTGGAG
GATCTCTCCAGCACTAGACGGATGAGCCTGGGTGCCCTGCTAGACTCTCA
25 CCAGCACTTGGCCGGTGCTGGCAGACGGCCCCACGCTTGCTGCTTAAAA
ACCTTCCTTAATAAAGCTGCAGTAGAAGCA

30. The nucleic acid of claim 27, which contains a nucleotidic sequence coding for at least part of the aminoacid sequence indicated hereafter :

GAGRODN

5 MetGlyAlaArgAsnSerValLeuArgGlyLysLysAlaAspGlu
 LeuGluArgIleArgLeuArgProGlyGlyLysLysLysTyrArg
 10 LeuLysHisIleValTrpAlaAlaAsnLysLeuAspArgPheGly
 100 LeuAlaGluSerLeuLeuGluSerLysGluGlyCysGlnLysIle
 LeuThrValLeuAspProMetValProThrGlySerGluAsnLeu
 15 LysSerLeuPheAsnThrValCysValIleTrpCysIleHisAla
 200 GluGluLysValLysAspThrGluGlyAlaLysGlnIleValArg
 300 ArgHisLeuValAlaGluThrGlyThrAlaGluLysMetProSer
 ThrSerArgProThrAlaProSerSerGluLysGlyGlyAsnTyr
 400 ProValGlnHisValGlyGlyAsnTyrThrHisIleProLeuSer
 25 ProArgThrLeuAsnAlaTrpValLysLeuValGluGluLysLys
 PheGlyAlaGluValValProGlyPheGlnAlaLeuSerGluGly
 300 CysThrProTyrAspIleAsnGlnMetLeuAsnCysValGlyAsp
 30 HisGlnAlaAlaMetGlnIleIleArgGluIleIleAsnGluGlu
 600 AlaAlaGluTrpAspValGlnHisProIleProGlyProLeuPro
 35 AlaGlyGlnLeuArgGluProArgGlySerAspIleAlaGlyThr
 700 ThrSerThrValGluGluGlnIleGlnTrpMetPheArgProGln

AsnProValProValGlyAsnIleTyrArgArgTrpIleGlnIle
 800
 GlyLeuGlnLysCysValArgMetTyrAsnProThrAsnIleLeu
 AspIleLysGlnGlyProLysGluProPheGlnSerTyrValAsp
 900
 ArgPheTyrLysSerLeuArgAlaGluGlnThrAspProAlaVal
 LysAsnTrpMetThrGlnThrLeuLeuValGlnAsnAlaAsnPro
 AspCysLysLeuValLeuLysGlyLeuGlyMetAsnProThrLeu
 1000
 GluGluMetLeuThrAlaCysGlnGlyValGlyGlyProGlyGln
 LysAlaArgLeuMetAlaGluAlaLeuLysGluValIleGlyPro
 1100
 AlaProIleProPheAlaAlaAlaGlnGlnArgLysAlaPheLys
 CysTrpAsnCysGlyLysGluGlyHisSerAlaArgGlnCysArg
 1200
 AlaProArgArgGlnGlyCysTrpLysCysGlyLysProGlyHis
 IleMetThrAsnCysProAspArgGlnAlaGlyPheLeuGlyLeu
 1300
 GlyProTrpGlyLysLysProArgAsnPheProValAlaGlnVal
 ProGlnGlyLeuThrProThrAlaProProValAspProAlaVal
 AspLeuLeuGluLysTyrMetGlnGlnGlyLysArgGlnArgGlu
 1400
 GlnArgGluArgProTyrLysGluValThrGluAspLeuLeuHis
 LeuGluGlnGlyGluThrProTyrArgGluProProThrGluAsp
 1500
 LeuLeuHisLeuAsnSerLeuPheGlyLysAspGln

31. The nucleic acid of claim 27, which contains a nucleotidic sequence coding for at least part of the aminoacid sequence indicated hereafter :

5 ArgLysAlaPheLys
 CysTrpAsnCysGlyLysGluGlyHisSerAlaArgGlnCysArg
 1200
 AlaProArgArgGlnGlyCysTrpLysCysGlyLysProGlyHis
 10 IleMetThrAsnCysProAspArgGlnAlaGlyPheLeuGlyLeu
 1300
 GlyProTrpGlyLysLysProArgAsnPheProValAlaGlnVal
 ProGlnGlyLeuThrProThrAlaProProValAspProAlaVal
 15 AspLeuLeuGluLysTyrMetGlnGlnGlyLysArgGlnArgGlu
 1400
 GlnArgGluArgProTyrLysGluValThrGluAspLeuLeuHis
 20 LeuGluGlnGlyGluThrProTyrArgGluProProThrGluAsp
 1500
 LeuLeuHisLeuAsnSerLeuPheGlyLysAspGln

32. The nucleic acid of claim 27, which contains a nucleotidic sequence coding for at least part of the aminoacid sequence indicated hereafter :

MetGlyAlaArgAsnSerValLeuArgGlyLysLysAlaAspGlu
5
LeuGluArgIleArgLeuArgProGlyGlyLysLysLysTyrArg
LeuLysHisIleValTrpAlaAlaAsnLysLeuAspArgPheGly
100
10 LeuAlaGluSerLeuLeuGluSerLysGluGlyCysGlnLysIle
LeuThrValLeuAspProMetValProThrGlySerGluAsnLeu
200
15 LysSerLeuPheAsnThrValCysValIleTrpCysIleHisAla
GluGluLysValLysAspThrGluGlyAlaLysGlnIleValArg
300
ArgHisLeuValAlaGluThrGlyThrAlaGluLysMetProSer
20 ThrSerArgProThrAlaProSerSerGluLysGlyGlyAsnTyr
400

33. The nucleic acid of claim 27, which contains a nucleotidic sequence coding for at least part of the aminoacid sequence indicated hereafter :

5 ProValGlnHisValGlyGlyAsnTyrThrHisIleProLeuSer
 ProArgThrLeuAsnAlaTrpValLysLeuValGluGluLysLys
 PheGlyAlaGluValValProGlyPheGlnAlaLeuSerGluGly
 500
 10 CysThrProTyrAspIleAsnGlnMetLeuAsnCysValGlyAsp
 HisGlnAlaAlaMetGlnIleIleArgGluIleIleAsnGluGlu
 600
 15 AlaAlaGluTrpAspValGlnHisProIleProGlyProLeuPro
 AlaGlyGlnLeuArgGluProArgGlySerAspIleAlaGlyThr
 700
 ThrSerThrValGluGluGlnIleGlnTrpMetPheArgProGln
 20 AsnProValProValGlyAsnIleTyrArgArgTrpIleGlnIle
 800
 GlyLeuGlnLysCysValArgMetTyrAsnProThrAsnIleLeu
 AspIleLysGlnGlyProLysGluProPheGlnSerTyrValAsp
 25 ArgPheTyrLysSerLeuArgAlaGluGlnThrAspProAlaVal
 900
 LysAsnTrpMetThrGlnThrLeuLeuValGlnAsnAlaAsnPro
 30 AspCysLysLeuValLeuLysGlyLeuGlyMetAsnProThrLeu
 1000
 GluGluMetLeuThrAlaCysGlnGlyValGlyGlyProGlyGln
 LysAlaArgLeuMetAlaGlnAlaLeuLysGluValIleGlyPro
 35 1100
 AlaProIleProPheAlaAlaAlaGlnGln

34. The nucleic acid of claim 27, which contains a nucleotidic sequence coding for at least part of the aminoacid sequence indicated hereafter :

ENVRM

5 MetMetAsnGlnLeuLeuIleAlaIleLeuLeuAlaSerAlaCys
 LeuValTyrCysThrGlnTyrValThrValPheTyrGlyValPro
 10 ThrTrpLysAsnAlaThrIleProLeuPheCysAlaThrArgAsn
 100 ArgAspThrTrpGlyThrIleGlnCysLeuProAspAspAsp
 TyrGlnGluIleThrLeuAsnValThrGluAlaPheAspAlaTrp
 15 AsnAsnThrValThrGluGlnAlaIleGluAspValTrpHisLeu
 200 PheGluThrSerIleLysProCysValLysLeuThrProLeuCys
 300 ValAlaMetLysCysSerSerThrGluSerSerThrGlyAsnAsn
 20 ThrThrSerLysSerThrSerThrThrThrThrThrProThrAsp
 400 GlnGluGlnGluIleSerGluAspThrProCysAlaArgAlaAsp
 25 AsnCysSerGlyLeuGlyGluGluGluThrIleAsnCysGlnPhe
 AsnMetThrGlyLeuGluArgAspLysLysLysGlnTyrAsnGlu
 300 ThrTrpTyrSerLysAspValValCysGluThrAsnAsnSerThr
 30 AsnGlnThrGlnCysTyrMetAsnHisCysAsnThrSerValIle
 600 ThrGluSerCysAspLysHisTyrTrpAspAlaIleArgPheArg
 35 TyrCysAlaProProGlyTyrAlaLeuLeuArgCysAsnAspThr
 700 AsnTyrSerGlyPheAlaProAsnCysSerLysValValAlaSer

ThrCysThrArgMetMetGluThrGlnThrSerThrTrpPheGly
 PheAsnGlyThrArgAlaGluAsnArgThrTyrIleTyrTrpHis
 GlyArgAspAsnArgThrIleIleSerLeuAsnLysTyrTyrAsn
 LeuSerLeuHisCysLysArgProGlyAsnLysThrVallysGln
 IleMetLeuMetSerGlyHisValPheHisSerHisTyrGlnPro
 IleAsnLysArgProArgGlnAlaTrpCysTrpPheLysGlyLys
 TrpLysAspAlaMetGlnGluValLysThrLeuAlaLysHisPro
 ArgTyrArgGlyThrAsnAspThrArgAsnIleSerPheAlaAla
 ProGlyLysGlySerAspProGluValAlaTyrMetTrpThrAsn
 CysArgGlyGluPheLeuTyrCysAsnMetThrTrpPheLeuAsn
 TrpIleGluAsnLysThrHisArgAsnTyrAlaProCysHisIle
 LysGlnIleIleAsnThrTrpHisLysValGlyArgAsnValTyr
 LeuProProArgGluGlyGluLeuSerCysAsnSerThrValThr
 SerIleIleAlaAsnIleAspTrpGlnAsnAsnAsnGlnThrAsn
 IleThrPheSerAlaGluValAlaGluLeuTyrArgLeuGluLeu
 GlyAspTyrLysLeuValGluIleThrProIleGlyPheAlaPro

ThrLysGluLysArgTyrSerSerAlaHisGlyArgHisThrArg

1500

GlyValPheValLeuGlyPheLeuGlyPheLeuAlaThrAlaGly

SerAlaMetGlyAlaArgAlaSerLeuThrValSerAlaGlnSer

1600

ArgThrLeuLeuAlaGlyIleValGlnGlnGlnGlnGlnLeuLeu

AspValValLysArgGlnGlnGluLeuLeuArgLeuThrValTrp

1700

GlyThrLysAsnLeuGlnAlaArgValThrAlaIleGluLysTyr

LeuGlnAspGlnAlaArgLeuAsnSerTrpGlyCysAlaPheArg

1800

GlnValCysHisThrThrValProTrpValAsnAspSerLeuAla

ProAspTrpAspAsnMetThrTrpGlnGluTrpGluLysGlnVal

ArgTyrLeuGluAlaAsnIleSerLysSerLeuGluGlnAlaGln

1900

IleGlnGlnGluLysAsnMetTyrGluLeuGlnLysLeuAsnSer

TrpAspIlePheGlyAsnTrpPheAspLeuThrSerTrpValLys

2000

TyrIleGlnTyrGlyValLeuIleIleValAlaValIleAlaLeu

ArgIleValIleTyrValValGlnMetLeuSerArgLeuArgLys

2100

GlyTyrArgProValPheSerSerProProGlyTyrIleGlnGln

IleEisIleEisLysAspArgGlyGlnProAlaAsnGluGluThr
 GluGluAspGlyGlySerAsnGlyGlyAspArgTyrTrpProTrp 2200
 ProIleAlaTyrIleHisPheLeuIleArgGlnLeuIleArgLeu
 LeuThrArgLeuTyrSerIleCysArgAspLeuLeuSerArgSer
 2300
 PheLeuThrLeuGlnLeuIleTyrGlnAsnLeuArgAspTrpLeu
 ArgLeuArgThrAlaPheLeuGlnTyrGlyCysGluTrpIleGln
 2400
 GluAlaPheGlnAlaAlaAlaArgAlaThrArgGluThrLeuAla
 GlyAlaCysArgGlyLeuTrpArgValLeuGluArgIleGlyArg
 2500
 GlyIleLeuAlaValProArgArgIleArgGlnGlyAlaGluIle
 AlaLeuLeu***GlyThrAlaValSerAlaGlyArgLeuTyrGlu
 2600
 TyrSerMetGluGlyProSerSerArgLysGlyGluLysPheVal
 GlnAlaThrLysTyrGly

35. The nucleic acid of anyone of claims 28 to 34 which is formed a recombinant nucleic acid comprising a nucleic acid from a vector and in which said cDNA or part of said cDNA is inserted.

5 36. The recombinant nucleic acid of claim 35 which is labelled.

37. A process for the detection of HIV-2 retrovirus or of its RNA in a biological liquid or tissue, particularly for the in vitro diagnosis in man of the potentiality or existence of LAS or of AIDS, which comprises contacting nucleic acids contained in said biological liquid or tissue with a probe containing a nucleic acid according to anyone of claims 28 to 36 under stringent hybridization conditions for the time necessary for said hybridization to occur, washing the hybride formed with a solution ensuring the preservation of said stringent conditions, and detecting the hybride formed.

38. A process for the production of HIV-2 retrovirus which comprises culturing human T4 lymphocytes or permanent cell lines derived from said T4 lymphocytes and carrying the T4 phenotype, which lymphocytes or cell lines had previously been infected with an isolate of HIV-2 virus and, particularly when the level of reverse transcriptase activity has reached a determined threshold, recovering and purifying the amounts of virus released in the culture medium of said lymphocytes or cell lines, particularly by differential centrifugation in a gradient of sucrose or metrizamide.

39. A process for the production of specific antigen of HIV-2 retrovirus which comprises lysing, particularly by means of detergent such as SDS (for instance 0.1 % SDS in a RIPA buffer) and recovering the lysate containing said antigens;

40. Process for the production of one of the

above defined proteins (p12, p16 or p26) or of a protein having the structure of gp140 or of determined parts of said proteins, which process comprises inserting the corresponding nucleic acid sequence in a vector capable of transforming an appropriate host, enabling the expression of an insert containing in said vector, transforming said host by said vector which comprises the said nucleotidic sequence, culturing the transformed cell lines host, recovering and purifying the expressed protein.

41. Process for the production of a hybridization probe for the detection of the RNA of HIV-2 retrovirus which comprises a DNA sequence, particularly according to anyone of claims 27 to 35, in a cloning vector by in vitro recombination, cloning the modified vector obtained in a competent cellular host, and recovering the DNA-recombinants obtained.

RR
G1

add
B1
add
C5
add
E1